Title:  Test-Retest Reliability of Measuring the Vertebral Arterial Blood Flow Velocity in People With Cervicogenic Dizziness

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Test-retest reliability of measuring the vertebral arterial blood flow velocity in people with cervicogenic dizziness.

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Abstract

Objectives

The purpose of this study was to determine the within session and between sessions reliability of measuring the vertebral artery blood flow velocities in people with cervicogenic dizziness using Doppler ultrasound at both upper and lower cervical levels.

Methods

Outcome measures were taken on two occasions three weeks apart with no active treatment provided in between the assessments on twelve participants. Pulsed-wave Doppler ultrasound was used to quantify time average mean velocities through the vertebral artery at upper cervical(C0-1) and lower cervical vertebrae (C5-6). The clinical outcome measures were also recorded in people with cervicogenic dizziness. Intraclass correlation coefficient was used to determine the within-session and between-session repeatability. Paired t-test was used to determine the differences in the time average mean velocities of blood flow at the same site of the vertebral artery and the clinical outcome measures in two sessions three weeks apart.

Results
In people with cervicogenic dizziness, there was no significant change in both clinical outcome measures and the time average mean velocities when the patients were measured three weeks apart (p > 0.05). This study demonstrated good within-session (ICC: 0.903 - 0.967) and between-session (ICC: 0.922 - 0.984) repeatability in measuring the vertical blood flow velocities in patients with cervicogenic dizziness when the clinical outcome measures were unchanged.

Conclusions

This study supports the use of Doppler ultrasound to identify changes in mean vertebral arterial blood flow velocities before and after intervention in people with cervicogenic dizziness in future studies.

*Key Words:* Doppler Ultrasound Imaging, Blood flow Velocity, Vertebral Artery, Dizziness, Reliability.
Introduction

Dizziness is a common problem that can lead to disability and impact on quality of life. In some cases of dizziness the cause can be attributed to pathology or dysfunction of upper cervical vertebral segments. This form of cervicogenic dizziness is characterized by symptoms of imbalance or spinning associated with neck pain, stiffness or headache.

It has been hypothesised that mechanical compression or stenosis of the vertebral artery could be one of the reasons leading to cervicogenic dizziness. Mechanical compression, tension, dissection, or stenosis of one of both vertebral arteries as they pass through the cervical region may reduce blood flow and thus result in symptoms of dizziness. Poor head and neck posture and mal-alignment of the upper cervical spine are among the causes of the mechanical compromise that could result in decreased vertebrobasilar blood flow velocity and lead to dizziness. However, to date no studies have investigated the test retest reliability in measuring the vertebral artery blood flow velocity in patients with cervicogenic dizziness over a period of time. It can be hypothesised that vertebral artery blood flow velocity would stay the same if the symptoms of cervicogenic dizziness stay the same. The results of this study provide evidence to support the use of Doppler ultrasound in investigating the role of vertebral arteries in management of cervicogenic dizziness in future studies.

Colour duplex/Doppler ultrasound is considered to provide a valid and reliable non-invasive measurement of vertebral arterial blood flow velocity. There are several studies reporting test-retest reliability of spectral Doppler ultrasound measures of vertebral artery blood flow. Schoning and Scheel reported same day repeat measure correlation coefficients in excess of 0.9 in measuring the cerebral time-averaged flow velocity and blood flow volume. Johnson et
al\textsuperscript{17} reported a good same session test-retest (intra-observer) intraclass correlation coefficient in the range of 0.80\,-\,0.94 for measures taken in left and right vertebral arteries velocities at the upper and lower cervical levels in asymptomatic participants. Previous work\textsuperscript{14,17} suggests that it would be feasible to measure the vertebral blood flow characteristics at the lower cervical level(C5-6) and atlanto-occipital (C0-1) level when the researchers adhere to protocol by taking account of potential confounding human factors such as consistency of gain settings, Doppler angle, and stabilisation of the ultrasound probe\textsuperscript{18}, and possible habituation effects\textsuperscript{16}.

There is evidence that Doppler ultrasound measures of vertebral artery blood flow may be sufficiently responsive to detect changes in cervical rotations\textsuperscript{13} or following intervention\textsuperscript{19}. It has been reported that a decrease in vertebral artery blood flow could be identified by Doppler ultrasound at both upper and lower cervical levels during end-range cervical rotation in asymptomatic participants\textsuperscript{12,20,21}. This demonstrated that Doppler ultrasound would be able to detect changes in the vertebral arterial blood flow velocities decreased when the vertebral arteries are under stress from compression or stretching. Doppler ultrasound has also been used to detect improvement in vertebral arterial blood flow velocities in patients with vertebrobasilar artery insufficiency before and after medical intervention\textsuperscript{19,22}. However, no studies have investigated upper and lower vertebral blood flow velocities in people with cervical dizziness. Consequently, it is unknown if there would be significant changes in vertebral artery blood flow velocities at the upper and lower cervical spine if clinical symptoms remain unchanged over a three week period in people with cervicogenic dizziness.
The aims of this study were to determine the within session and between sessions reliability of measuring the vertebral artery blood flow velocities in people with cervicogenic dizziness using Doppler ultrasound at both upper and lower cervical levels. It is hypothesized that no significant changes would occur in the vertebral artery velocities if there are no significant changes in the clinical symptoms in people with cervicogenic dizziness. The results of this study provide a basis to investigate the role of upper and lower vertebral arterial blood flow velocities and to support the use of Doppler ultrasound in determining the effect of intervention on the blood flow velocities in the management of cervicogenic dizziness in further studies.
Methods

Participants

The experimental protocol was approved by the Faculty Research Ethics Committee of Faculty of Health, Education and Society, Plymouth University, United Kingdom and informed consent for the study was obtained from all participants per the WORLD Medical Association Declaration of Helsinki: Ethical principles for medical research involving human subjects, 2008. The sample size was estimated with an effect size of 0.75, a power of 0.80 and a significance level of 0.05. A sample size of twelve was deemed to be appropriate for this matched pair study design. Therefore, twelve participants with symptoms of dizziness and neck pain as their main complaint were recruited with the following inclusion criteria:

- Dizziness described as imbalance or unsteadiness.
- Dizziness related to either movements or positions of the cervical spine, or occurring with a stiff or painful neck.
- Symptoms lasting for at least 4 weeks.
- 18-55 years of age.

Participants were excluded if they had:

- Previous history of stroke, or any neurological disorders.
- Chiropractic care or physiotherapy within the past 4 weeks.
- Currently receiving treatment for dizziness/neck pain by other healthcare providers.
- Presence of inflammatory joint disease, infection, tumour, or fracture of the spine or cranium, central vascular/ neurologic condition suspected of causing neck pain and/or dizziness/vertigo or other conditions contraindicating high-velocity, small-amplitude spinal manipulative therapy.
- Evidence of narcotic or other drug abuse.
- An on-going personal injury or workers' compensation case related to dizziness/vertigo or neck pain or currently seeking or receiving disability for dizziness/vertigo or neck pain.

The baseline characteristics of the participants are presented in Table 1.

Equipment and procedures

Outcome measures were taken on two occasions three weeks apart with no active treatment provided in between the assessments. Both the Doppler ultrasound readings and clinical outcome measures were taken by the same examiner.

Pulsed-wave Doppler ultrasound (Model: MySonoU5, Medison Co., Ltd, Seoul, South Korea) was used to quantify laminar blood flow velocity through the vertebral artery at upper (C0-1) and lower cervical vertebrae (C5-6)\textsuperscript{13,15}. All blood flow measurements were taken in a quiet, dimmed environment with the patient supine with head supported in neutral (no flexion or extension) and turned 10 degrees to the contralateral side to allow ease of probe placement and optimal imaging of the artery at both upper and lower level. The measurements were taken in the order of lower right, lower left, upper right, upper left at the cervical spine for each participant. An initial image was taken at each site followed by a series of three measurements to ensure reliability of the measurement\textsuperscript{13,17,20}. The Doppler sample gate was set at its smallest size and placed in the center of the vessel in order to standardize the signal received.
The lower vertebral artery measurement was taken at C5-6 level which is typically the first entry level of the vertebral artery into the transverse foramen of the cervical spine (Figure 1). The transducer was placed longitudinally at the anterolateral aspect of the neck and the probe shifted laterally until the cervical transverse processes were visualised and the entry of the vertebral artery into the transverse foramen identified. The vertebral artery was then tracked upwards and the measurement taken between the transverse processes of the C5-6 (between fifth cervical vertebra (C5) and sixth cervical vertebra (C6). The upper level measurement was taken at atlanto-occipital (C0-1) as the vertebral artery exited from the transverse foramen of atlas (C1) and passed onto the posterior arch of the atlas (Figure 2). The probe was placed transversely below the mastoid process, posterior to the angle of the jaw.

Once optimal imaging of the artery at both upper and lower cervical level have been acquired, off-line analysis was carried by a separate researcher. The time average mean velocities were determined with the angle of insonation adjusted at 60 degrees (Figures 1 and 2). The order of the ultrasound imaging analysis has been randomised and the researcher was blinded to the order of the Doppler ultrasound imaging and clinical outcome measures of the participants.

Clinical outcome measures

Disability caused by dizziness was measured with the Dizziness Handicap Inventory. The Dizziness Handicap Inventory assesses the impact of dizziness on the functional, emotional and physical aspects of everyday life. The highest possible score is 100, indicating maximum self-perceived handicap. The Dizziness Handicap Inventory has been shown to be
a highly reliable and responsive tool.\textsuperscript{26,27} Significant correlations between Dizziness Handicap Inventory scores and specific objective measures of balance and gait have been demonstrated.\textsuperscript{28}

Dynamic balance in walking was measured by the Dynamic Gait Index\textsuperscript{29}. Dynamic Gait Index assessed dynamic postural stability as functional gait according to eight tasks with varying demands including walking at different speeds, walking with head movements, negotiating obstacles, turning, ascending and descending stairs. Each item was rated on a 4-level ordinal scale, with a maximum score of 24\textsuperscript{29}. The Dynamic Gait Index has been regularly used as a outcome measures in clinical trial for patients with balance problems\textsuperscript{30,31}.

Severity of dizziness (an average level over the previous few days) was measured with a 10 cm visual analogue scale. The visual analogue scale has been used to measure dizziness in other clinical studies.\textsuperscript{5}

Frequency of dizziness was measured on a six-point rating scale (0=no dizziness, 1=dizziness less than once per month, 2=1–4 episodes of dizziness per month, 3=1–4 episodes of dizziness per week, 4=dizziness once daily, 5=dizziness more than once a day or constant).\textsuperscript{32}

Active cervical range of motion was measured with a strap-on head goniometer consisting of an inclinometer dial for measuring flexion, extension and lateral flexion and a compass dial for measuring rotation (OB ‘‘Myrin’’ goniometer, OB Rehab, Solna, Sweden).\textsuperscript{33}

Statistical analysis
Statistical analysis was carried out with SPSS software (Version 19.0). Intraclass correlation coefficient (ICC$_{3,k}$) with 95% confidence interval and standard error of measurement (SEM) was calculated to determine intra-rater reliability of the three repeat measures of time average mean velocities of the upper and lower vertebral arterial blood flow in each session.

Paired t-test was used to determine the differences in the time average mean velocities of blood flow at the same site of the vertebral artery and the clinical outcome measures in two sessions three weeks apart. Level of significance was set at 0.05. Post hoc power calculation was carried out to determine the type II error when no significant differences were found in the outcome measures between the two sessions.
Results

Participant characteristics were presented in table 1.

The mean ICC\textsubscript{3,k} for measuring the blood flow velocities in the vertebral artery at upper and lower cervical levels were found to be highly repeatable within each session (Table 2 and 3) and between session (Table 4). All the ICC values were greater than 0.900. The ICC ranges from 0.903 - 0.957 in the first session, 0.930 - 0.967 in the second session and 0.922 - 0.984 between the two sessions three weeks apart.

Paired t-tests revealed no significant statistical differences in both the time average mean velocities and clinical outcome measures between the two assessment sessions three weeks apart (p > 0.05, Table 5 and 6).

The clinical outcome measures suggested that patients with cervicogenic dizziness in this study were being affected significantly in the balance and quality of life. There was a high pain level, severity of dizziness and frequency of dizziness (Table 6). The mean score of 48.11 out of 100 of the Dizziness Handicap Inventory suggested that this group of patients with cervicogenic dizziness have moderate handicap\textsuperscript{25} (Table 6).
Discussion

This is the first study to report on a method both to measure vertebral arterial blood flow velocity by Doppler ultrasound in patients with cervicogenic dizziness and to investigate its reliability over a three week period. The reliability of the described method was found to be excellent within sessions and between sessions. The technique would be highly repeatable and sufficiently precise to support its use as a research tool and in clinical practice.35

Doppler ultrasound has been reported to be sensitive enough to detect changes in vertebral arterial blood flow parameters in patients with vertebrobasilar artery insufficiency before and after medical intervention 19,22. In addition, Doppler ultrasound has also been used to identify blood flow changes following at the end range of motion of neck rotation in asymptomatic participants, which could potentially exert pressure on the vertebral artery and reduce the blood flow velocity.12,20,21

In patients with cervicogenic dizziness and moderate handicap, there was no significant change in both clinical outcome measures and the time average mean velocities when the patients were measured three weeks apart (p > 0.05, Table 5 and 6). A compromise post-hoc power analysis was carried out to identify the statistical power 36 37 of the Doppler vertebral blood flow assessments, vertebral arterial blood flow, the statistical power was excellent, ranging from 88.59% to 99.99%, ruling out the chance of type II error (false-negative) (Table 5).
This study demonstrated an excellent between-session repeatability (ICC: 0.922 - 0.984) in measuring the vertebral arterial blood flow velocities in patients with cervicogenic dizziness. The vertebral arterial blood flow is readily identifiable at the upper and lower cervical spine; the use of body landmarks and the real time visualisation of the entry of the vertebral artery to transverse foramen from the Doppler imaging as well as standardisation of protocol described in the methodology ensured good repeatability.

Limitations

There were some individual variations in the blood flow velocities with quite a large standard deviation (Table 5). This may limit the use of Doppler ultrasound as a diagnostic tool in assessing patients with cervicogenic dizziness. However, we have demonstrated that determination of time average mean velocities from the Doppler ultrasound at the upper and lower vertebral arteries could be used for evaluation for within-subject pairwise comparison in clinical population. In this study, one of the limitations is that the order of measuring was not randomized which was to ensure the standardization of the procedures.

Conclusion

This study provides a foundation for the use of Doppler ultrasound in the assessment of patients with cervicogenic dizziness. The repeatability of time average mean velocities was excellent between measurements taken three weeks apart when there was no change in clinical outcome measures. Therefore, measuring vertebral arterial blood flow velocities at C0-1 and C5-6 could provide a clinically meaningful outcome measure to identify potential
Changes in mean blood flow velocity before and after intervention in people with cervicogenic dizziness in future studies. If the clinical outcome of patients improved after an intervention, it would be valuable to see if changes occur in the time averaged blood flow velocities as this would demonstrate the role of vertebral artery blood flow in patients with cervicogenic dizziness.

This is the first study to measure the blood flow velocities in patients with cervicogenic dizziness over a three week period. The Doppler ultrasound findings demonstrated a very good reliability and it corroborated with the clinical outcome measurements. Further research is planned to provide an intervention to improve the symptoms in people with cervicogenic dizziness and investigate if blood flow changes will occur when the clinical symptoms improve. This will improve our understanding of the role of vertebral artery involvement in cervicogenic dizziness.

Acknowledgment

This study was funded by Australian Spinal Research Foundation Research Grant LG2011-4.
References


**Figure legends**

Figure 1. Doppler ultrasound imaging measured at the lower cervical spine between fifth cervical vertebra and sixth cervical vertebra.
Figure 2. Doppler ultrasound imaging measured at the upper cervical spine at atlanto-occipital joint as the vertebral artery exited from the transverse foramen of atlas and passed onto the posterior arch of the atlas.
Table 1. Baseline characteristics of participants

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td>45.1 ± 9.2</td>
</tr>
<tr>
<td>Weight (kg)</td>
<td>72.64 ± 10.80</td>
</tr>
<tr>
<td>Height (m)</td>
<td>1.68 ± 0.07</td>
</tr>
<tr>
<td>Onset of symptoms (months)</td>
<td>94.0 ± 65.9</td>
</tr>
<tr>
<td>Gender</td>
<td>Male: 5; Female: 7</td>
</tr>
</tbody>
</table>
Table 2. Within-session Intraclass correlation (ICC) at the first session

<table>
<thead>
<tr>
<th></th>
<th>ICC of the time averaged mean velocities (TAMV)</th>
<th>95% Confidence Interval (CI)</th>
<th>Standard Error of Measurement (SEM)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Right vertebral artery at C5/6</td>
<td>0.914</td>
<td>0.805-0.965</td>
<td>1.05</td>
</tr>
<tr>
<td>Left vertebral artery at C5/6</td>
<td>0.957</td>
<td>0.916-0.980</td>
<td>0.99</td>
</tr>
<tr>
<td>Right vertebral artery at C0/1</td>
<td>0.903</td>
<td>0.789-0.965</td>
<td>1.11</td>
</tr>
<tr>
<td>Left vertebral artery at C0/1</td>
<td>0.908</td>
<td>0.783-0.962</td>
<td>1.30</td>
</tr>
</tbody>
</table>
Table 3. Within-session Intraclass correlation (ICC) at the second session (three weeks later)

<table>
<thead>
<tr>
<th>Artery Location</th>
<th>ICC of the time averaged mean velocities (TAMV)</th>
<th>95% Confidence Interval (CI)</th>
<th>Standard Error of Measurement (SEM)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Right vertebral artery at C5/6</td>
<td>0.930</td>
<td>0.864-0.967</td>
<td>0.98</td>
</tr>
<tr>
<td>Left vertebral artery at C5/6</td>
<td>0.965</td>
<td>0.933-0.984</td>
<td>0.85</td>
</tr>
<tr>
<td>Right vertebral artery at C0/1</td>
<td>0.967</td>
<td>0.936-0.984</td>
<td>0.63</td>
</tr>
<tr>
<td>Left vertebral artery at C0/1</td>
<td>0.937</td>
<td>0.878-0.970</td>
<td>1.18</td>
</tr>
</tbody>
</table>
Table 4. Intraclass correlation (ICC) between the two sessions (first session vs second session)

<table>
<thead>
<tr>
<th></th>
<th>ICC of the time averaged mean velocities (TAMV)</th>
<th>95% Confidence Interval (CI)</th>
<th>Standard Error of Measurement (SEM)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Right vertebral artery at C5/6</td>
<td>0.984</td>
<td>0.944-0.995</td>
<td>0.46</td>
</tr>
<tr>
<td>Left vertebral artery at C5/6</td>
<td>0.974</td>
<td>0.911-0.993</td>
<td>0.75</td>
</tr>
<tr>
<td>Right vertebral artery at C0/1</td>
<td>0.968</td>
<td>0.889-0.991</td>
<td>0.63</td>
</tr>
<tr>
<td>Left vertebral artery at C0/1</td>
<td>0.922</td>
<td>0.871-0.959</td>
<td>1.25</td>
</tr>
</tbody>
</table>
Table 5. Time averaged mean velocities (TAMV) of the two sessions

<table>
<thead>
<tr>
<th></th>
<th>TAMV (cm/s) (mean ± SD)</th>
<th>P</th>
<th>Power (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>First session</td>
<td>Second session (3 weeks later)</td>
<td></td>
</tr>
<tr>
<td>Right vertebral artery at C5/6</td>
<td>14.07 ± 3.57</td>
<td>14.28 ± 3.69</td>
<td>0.431</td>
</tr>
<tr>
<td>Left vertebral artery at C5/6</td>
<td>12.51 ± 4.78</td>
<td>12.06 ± 4.57</td>
<td>0.316</td>
</tr>
<tr>
<td>Right vertebral artery at C0/1</td>
<td>11.17 ± 3.58</td>
<td>10.93 ± 3.47</td>
<td>0.500</td>
</tr>
<tr>
<td>Left vertebral artery at C0/1</td>
<td>13.73 ± 4.29</td>
<td>13.63 ± 4.69</td>
<td>0.943</td>
</tr>
</tbody>
</table>

Paired t test revealed no significant differences in the time averaged mean velocities (cm/s) between two sessions three weeks apart.
Table 6. Clinical outcome measures between the two sessions

<table>
<thead>
<tr>
<th>Outcome Measure</th>
<th>First session</th>
<th>Second session (3 weeks later)</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Dizziness Handicap Inventory (Max: 100)</td>
<td>48.11 ± 16.19</td>
<td>48.15 ± 15.33</td>
<td>0.957</td>
</tr>
<tr>
<td>Dynamic gait index (Max: 24)</td>
<td>21.69 ± 1.55</td>
<td>22.15 ± 1.46</td>
<td>0.351</td>
</tr>
<tr>
<td>Visual analogue scale of neck pain (Max: 10)</td>
<td>7.31 ± 1.7</td>
<td>6.69 ± 1.65</td>
<td>0.359</td>
</tr>
<tr>
<td>Visual analogue scale of severity of dizziness (Max: 10)</td>
<td>5.69 ± 1.75</td>
<td>5.84 ± 1.68</td>
<td>0.165</td>
</tr>
<tr>
<td>Frequency of dizziness (Max: 5)</td>
<td>3.85 ± 1.21</td>
<td>3.77 ± 1.42</td>
<td>0.584</td>
</tr>
<tr>
<td>Neck range of motion (degrees)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Flexion</td>
<td>38.08 ± 10.9</td>
<td>41.92 ± 9.9</td>
<td>0.192</td>
</tr>
<tr>
<td>Extension</td>
<td>41.15 ± 9.61</td>
<td>42.69 ± 8.81</td>
<td>0.165</td>
</tr>
<tr>
<td>Rotation at the more restricted side</td>
<td>53.85 ± 11.21</td>
<td>51.54 ± 9.21</td>
<td>0.165</td>
</tr>
<tr>
<td>Rotation at the less restricted side</td>
<td>60.00 ± 10.41</td>
<td>56.92 ± 9.69</td>
<td>0.431</td>
</tr>
<tr>
<td>Side flexion at the more restricted side</td>
<td>28.08 ± 6.63</td>
<td>30.00 ± 7.91</td>
<td>0.175</td>
</tr>
<tr>
<td>Side flexion at the less restricted side</td>
<td>31.92 ± 6.30</td>
<td>30.38 ± 8.53</td>
<td>0.264</td>
</tr>
</tbody>
</table>

Paired t test revealed no significant differences in the clinical outcome measures (p > 0.05).